

Pending Claims as of November 16, 2001

76. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

77. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

78. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

79. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

80. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCATTTCTCAATGGGACGGAGCGGGTGCGGTTCTTGGAGA  
GACACTTCCATAACCAGGAGGAGTACGCGCGCTTCGACAGCG  
ACGTGGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACCTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCTGGTGATGCTAGAAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGGCTTTGTGCTGGGCCTGCTC  
TTCCTTGGGGCCGGGCTGTTTCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;
- (ii) GGGGACACCCGACCACGTTTCTTGGAGCAGGTAAACATGAG  
TGTCATTTCTTCAACGGGACGGAGCGGGTGCGGTTCTTGGAC  
AGATACTTCTATACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACCTACGGG  
GTTGGTGAGAGCTTCACAGTGCAGCGGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCTGGTGATGCTGGAAACAGTTCTCGGAGTGGA  
GAGGTTTACACCTCCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGGCTTCGTGCTGGGCCTG  
CTCTTCTTGGGGCCGGGCTGTTTCTACTTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;
- (iii) a DNA sequence which is fully  
complementary to the DNA sequence of (i)  
or (ii); and
- (iv) a DNA sequence which differs from the DNA  
sequence of (i) or (ii) in codon sequence  
due to the degeneracy of the genetic code,  
and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

81. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCATTTCTCAATGGGACGGAGCGGGTGCGGTTCTGGAGA  
GACACTTCCATAAACCAGGAGGAGTACGCGCGCTTCGACAGCG  
ACGTGGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACCTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCGAGTCCATCCTCAGG  
TGAAGTGTGTATCCTGCAAGACCCAGCCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCTGGTGTATGCTAGAAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGGCTTTGTGCTGGGCCTGCTC  
TTCCTTGGGGCCGGGCTGTTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;
- (ii) GGGGACACCCGACCACGTTTCTTGGAGCAGGTTAAACATGAG  
TGTCATTTCTTCAACGGGACGGAGCGGGTGCGGTTCTGGAC  
AGATACTTCTATACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACCTACGGG  
GTTGGTGAGAGCTTCACAGTGCAGCGGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCTGGTGTATGCTGGAAACAGTTCTCCTCGGAGTGGA  
GAGGTTTACACCTCCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGGAAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGGCTTCGTGCTGGGCCTG  
CTCTTCCTTGGGGCCGGGCTGTTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;
- (iii) a DNA sequence which is fully  
complementary to the DNA sequence of (i)  
or (ii); and
- (iv) a DNA sequence which differs from the DNA  
sequence of (i) or (ii) in codon sequence  
due to the degeneracy of the genetic code,  
and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and
- (b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

83. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and
- (d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

84. An HLA-DR typing process comprising the steps of:

- (a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and
- (b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

85. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;  
(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(d) detecting areas of hybridization between said DNA to be typed and said second DNA.

86. The HLA-DR typing process according to claim 76 or 78, wherein said DNA sequence is characterized by a nucleotide sequence selected from the group consisting of:

TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA, GGGGCCAGGTGGACAATTA, TGGAGCAGGTAAACATGA, TCCTGGACAGATACTTCTA and GGGCCGCGGTGGACACCTA.

87. The HLA-DR typing process according to claim 77 or 79, wherein said second DNA is characterized by a nucleotide sequence selected from the group consisting of:

TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA, GGGGCCAGGTGGACAATTA, TGGAGCAGGTAAACATGA, TCCTGGACAGATACTTCTA and GGGCCGCGGTGGACACCTA.

88. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said DNA sequence.

89. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said second DNA.

90. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said DNA in said sample to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

91. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said size-fractionated DNA

to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

92. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein said DNA sequence is a labeled DNA sequence and its label is used for detecting hybridization between said DNA in said sample and said DNA sequence.

93. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein said second DNA is a labeled DNA and its label is used for detecting hybridization between said size-fractionated DNA and said second DNA.

94. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (ii) DNA sequences encoding amino acids 26-32 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iii) DNA sequences encoding amino acids 72-78 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

95. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

96. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of

hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences.

97. The HLA-DR typing kit according to any one of claims 94, 95 or 96, wherein said DNA sequence is labeled.

98. The HLA-DR typing kit according to any one of claims 94, 95 or 96, further comprising a 19-mer hybridization control, said hybridization control being a DNA being the nucleotide sequence: GCTTCGACAGCGACGTGGG.

99. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of an HLA-DR- $\beta$  locus of the human lymphocyte antigen complex, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

100. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

101. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting



- essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C, and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences.

102. The HLA-DR typing kit according to anyone of claims 99, 100 or 101, wherein said DNA sequence is labeled.

Pending Claims as of April 2002

76. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

77. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

78. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

79. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

80. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTTCTTGAGCTGCTTAAGTCTGAG  
TGTCATTTCTCAATGGGACGGAGCGGGTGCGGTTCTGGAGA  
GACACTTCCATAACCAGGAGGAGTACGCGCGCTTCGACAGCG  
ACGTGGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACCTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCGAGTCCATCCTCAGG  
TGAAGTCACTGCTGCTGTGAGTGGTTTCTATCCAGGCAGCAT  
TGAAGTCACTGCTGCTGTGAGTGGTTTCTATCCAGGCAGCAT  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCTGGTGTGCTAGAAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGGCTTTGTGCTGGGCCTGCTC  
TTCCTTGGGGCCGGGCTGTTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;  
(ii) GGGGACACCCGACCACGTTTCTTGAGCAGGTTAAACATGAG  
TGTCATTTCTTCAACGGGACGGAGCGGGTGCGGTTCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACCTACGGG  
GTTGGTGTGAGAGCTTCACAGTGCAGCGGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCTGGTGTGCTGGAAACAGTTCCTCGGAGTGGAG  
GAGGTTTACACCTCCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGGAAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGGCTTCGTGCTGGGCCTG  
CTCTTCCTTGGGGCCGGGCTGTTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTCTGAGC;  
(iii) a DNA sequence which is fully  
complementary to the DNA sequence of (i)  
or (ii); and  
(iv) a DNA sequence which differs from the DNA  
sequence of (i) or (ii) in codon sequence  
due to the degeneracy of the genetic code,  
and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

81. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCATTTTCTCAATGGGACGGAGCGGGTGCGGTTCCTGGAGA  
GACACTTCCATAACCAGGAGGAGTACGCGCGCTTCGACAGCG  
ACGTGGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACCTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCTGGTGTATGCTAGAAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGGCTTTGTGCTGGGCCTGCTC  
TTCCTTGGGGCCGGGCTGTTTCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCCTGAGC;
- (ii) GGGGACACCCGACCACGTTTCTTGGAGCAGGTAAACATGAG  
TGTCATTTTCTTCAACGGGACGGAGCGGGTGCGGTTCCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACCTACGGG  
GTTGGTGTGAGAGCTTCACAGTGCAGCGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCTGGTGTATGCTGGAAACAGTTCTCGGAGTGGAG  
GAGGTTTACACCTCCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGGCTTCGTGCTGGGCCTG  
CTCTTCTTGGGGCCGGGCTGTTTCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCCTGAGC;
- (iii) a DNA sequence which is fully  
complementary to the DNA sequence of (i)  
or (ii); and
- (iv) a DNA sequence which differs from the DNA  
sequence of (i) or (ii) in codon sequence  
due to the degeneracy of the genetic code,  
and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCATTTCTCAATGGGACGGAGCGGGTGCGGTTCTGGAGA  
GACACTTCCATAACCAGGAGGAGTACGCGCGCTTCGACAGCG  
ACGTGGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACCTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCTGGTGATGCTAGAAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGGCTTTGTGCTGGGCCTGCTC  
TTCCTTGGGGCCGGGCTGTTTCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCCTGAGC;
- (ii) GGGGACACCCGACCACGTTTCTTGGAGCAGGTTAAACATGAG  
TGTCATTTCTTCAACGGGACGGAGCGGGTGCGGTTCTGGAC  
AGATACTTCTATACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACCTACGGG  
GTTGGTGAGAGCTTCACAGTGCAGCGGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAGACCCAGCCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCTGGTGATGCTGGAAACAGTTTCTCGGAGTGGGA  
GAGGTTTACACCTCCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGGCTTCGTGCTGGGCCTG  
CTCTTCTTGGGGCCGGGCTGTTTCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCCTGAGC;
- (iii) a DNA sequence which is fully  
complementary to the DNA sequence of (i)  
or (ii); and
- (iv) a DNA sequence which differs from the DNA  
sequence of (i) or (ii) in codon sequence  
due to the degeneracy of the genetic code,  
and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and
- (b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

83. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and
- (d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

84. An HLA-DR typing process comprising the steps of:

- (a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and
- (b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

85. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;  
(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and
- (d) detecting areas of hybridization between said DNA to be typed and said second DNA.

86. The HLA-DR typing process according to claim 76 or 78, wherein said DNA sequence is characterized by a nucleotide sequence selected from the group consisting of:

TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA,  
GGGGCCAGGTGGACAATTA, TGGAGCAGGTAAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

87. The HLA-DR typing process according to claim 77 or 79, wherein said second DNA is characterized by a nucleotide sequence selected from the group consisting of:  
TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA, GGGGCCAGGTGGACAATTA,  
TGGAGCAGGTAAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

88. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said DNA sequence.

89. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said second DNA.

90. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said DNA in said sample to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

91. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said size-fractionated DNA



to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

92. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein said DNA sequence is a labeled DNA sequence and its label is used for detecting hybridization between said DNA in said sample and said DNA sequence.

93. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein said second DNA is a labeled DNA and its label is used for detecting hybridization between said size-fractionated DNA and said second DNA.

94. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (ii) DNA sequences encoding amino acids 26-32 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iii) DNA sequences encoding amino acids 72-78 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

95. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

96. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of

hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences.

97. The HLA-DR typing kit according to any one of claims 94, 95 or 96, wherein said DNA sequence is labeled.

98. The HLA-DR typing kit according to any one of claims 94, 95 or 96, further comprising a 19-mer hybridization control, said hybridization control being a DNA being the nucleotide sequence: GCTTCGACAGCGACGTGGG.

99. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of an HLA-DR- $\beta$  locus of the human lymphocyte antigen complex, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

100. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

101. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting

- essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C, and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences.

102. The HLA-DR typing kit according to anyone of claims 99, 100 or 101, wherein said DNA sequence is labeled.

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27. An isolated DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing, at to a polymorphic region of said locus to allow determination of one or more HLA alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

(a) DNA sequences encoding amino acids 8-14 of said locus;

(b) DNA sequences encoding amino acids 26-32 of said locus;

(c) DNA sequences encoding amino acids 72-78 of said locus;

(d) DNA sequences which are portions of any one of the foregoing DNA sequences and which are capable of hybridizing to said polymorphic region;

(e) DNA sequences which differ from any of the foregoing DNA sequences in codon sequence due to the degeneracy of the genetic code;

(f) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and

(g) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

29. An isolated DNA sequence encoding a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being selected from the group consisting of:

(a) DNA sequences encoding amino acids 8-14 of said locus;

(b) DNA sequences encoding amino acids 26-32 of said locus;

(c) DNA sequences encoding amino acids 72-78 of said locus;

(d) DNA sequences which are portions of any one of the foregoing DNA sequences and which are capable of hybridizing to said polymorphic region;

(e) DNA sequences which differ from any of the foregoing DNA sequences in codon sequence due to the degeneracy of the genetic code; and

(f) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

33. An isolated DNA sequence selected from the group consisting of:

(a) DNA sequences encoding a majority of the region defined by amino acids:

- (i) 8-14,
- (ii) 26-32,
- (iii) 39-45, or
- (iv) 72-78

of the polypeptide coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;

(b) DNA sequences that are allelic variants of any of the foregoing DNA sequences; and

(c) DNA sequences that are complementary to any of the foregoing DNA sequences.